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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/693,802	10/23/2003	Ivo Franci Eggen	O 2000.662 USD2	2360
31846 7590 01/25/2007 INTERVET INC. PATENT DEPARTMENT PO BOX 318 MILLSBORO, DE 19966-0318			EXAMINER EPPERSON, JON D	
			ART UNIT	PAPER NUMBER
			1639	
SHORTENED STATUTORY PERIOD OF RESPONSE		MAIL DATE	DELIVERY MODE	
3 MONTHS		01/25/2007	PAPER	

**Please find below and/or attached an Office communication concerning this application or proceeding.**

If NO period for reply is specified above, the maximum statutory period will apply and will expire 6 MONTHS from the mailing date of this communication.

**Office Action Summary**

Application No.

10/693,802

Applicant(s)

EGGEN ET AL.

Examiner

Jon D. Epperson

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-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

**Period for Reply**

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) OR THIRTY (30) DAYS, WHICHEVER IS LONGER, FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

**Status**

- 1) ☒ Responsive to communication(s) filed on 10 October 2006.
- 2a) ☐ This action is FINAL. 2b) ☒ This action is non-final.
- 3) ☐ Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

**Disposition of Claims**

- 4) ☒ Claim(s) 28-46 and 48-55 is/are pending in the application.
- 4a) Of the above claim(s) 40, 43 and 50-54 is/are withdrawn from consideration.
- 5) ☐ Claim(s) \_\_\_\_\_ is/are allowed.
- 6) ☒ Claim(s) 28-39, 41, 42, 44-46, 48, 49 and 55 is/are rejected.
- 7) ☐ Claim(s) \_\_\_\_\_ is/are objected to.
- 8) ☐ Claim(s) \_\_\_\_\_ are subject to restriction and/or election requirement.

**Application Papers**

- 9) ☐ The specification is objected to by the Examiner.
- 10) ☐ The drawing(s) filed on \_\_\_\_\_ is/are: a) ☐ accepted or b) ☐ objected to by the Examiner.  
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).  
Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
- 11) ☐ The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

**Priority under 35 U.S.C. § 119**

- 12) ☐ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
- a) ☐ All b) ☐ Some \* c) ☐ None of:
1. ☐ Certified copies of the priority documents have been received.
  2. ☐ Certified copies of the priority documents have been received in Application No. \_\_\_\_\_.
  3. ☐ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

\* See the attached detailed Office action for a list of the certified copies not received.

**Attachment(s)**

- |  |   |
|--|---|
| 1) <input checked="" type="checkbox"/> Notice of References Cited (PTO-892)            | 4) <input type="checkbox"/> Interview Summary (PTO-413)           |
| 2) <input type="checkbox"/> Notice of Draftsperson's Patent Drawing Review (PTO-948)   | Paper No(s)/Mail Date. _____                                      |
| 3) <input checked="" type="checkbox"/> Information Disclosure Statement(s) (PTO/SB/08) | 5) <input type="checkbox"/> Notice of Informal Patent Application |
| Paper No(s)/Mail Date <u>10/10/06</u> .  | 6) <input type="checkbox"/> Other: _____                          |

## DETAILED ACTION

### *Request for Continued Examination (RCE)*

1. A request for continued examination (RCE) under 37 CFR 1.114, including the fee set forth in 37 CFR 1.17(e), was filed in this application after final rejection. Since this application is eligible for continued examination under 37 CFR 1.114, and the fee set forth in 37 CFR 1.17(e) has been timely paid, the finality of the previous Office action has been withdrawn pursuant to 37 CFR 1.114. Applicant's submission filed on 10/10/06 has been entered. Claims 28-46 and 48-55 were pending. Applicants amended claims 28, 29 and 48. No claims were added or canceled. Therefore, claims 28-46 and 48-55 are currently pending. Claims 40, 43 and 50-54 are drawn to non-elected species and/or inventions and thus these claims remain withdrawn from further consideration by the examiner, 37 CFR 1.142(b), there being no allowable generic claim. Therefore, claims 28-39, 41, 42, 44-46, 48, 49 and 55 are examined on the merits.

Those sections of Title 35, US code, not included in the instant action can be found in previous office actions.

### *IDS*

2. The information disclosure statement filed October 10, 2000, fails, in part, to comply with the provisions of 37 CFR 1.97, 1.98 and MPEP § 609 because one publication cited therein, Eggen, I.F. et al. "Rapid solution-phase synthesis ...", lack a publication date, a necessary element for consideration. While the other patent and other publications cited therein, and supplied, therewith, have been considered as to the merits, the above cited publications have not.

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Applicant is advised that the date of any re-submission of these citations contained in this information disclosure statement or the submission of the missing element – their publication dates – will be the date of submission for purposes of determining compliance with the requirements based on the time of filing the statement, including all certification requirements for statements under 37 CFR 1.97(e). See MPEP § 609 C(1).

### **Withdrawn Objections/Rejections**

3. The objection to claim 28 is withdrawn in view of Applicants' amendments thereto (i.e., removal of extra "and"). The statutory double patenting rejection over claims 53 and 54 is hereby withdrawn in view of Applicants' arguments (e.g., see 10/10/06, Response, top of page 7 noting that these claims are withdrawn). The Examiner regrets the inadvertent error. All other rejections are maintained and the arguments are addressed below.

### **Outstanding Objections and/or Rejections**

#### ***Claim Rejections - 35 USC § 102***

4. Claims 28, 30, 31, 36, 41, 42, 44-46, 49 and 55 are rejected under 35 U.S.C. 102(b) as being anticipated by Carpino et al. (Carpino, et al. "The 1,1-Dioxobenzo[b]thiophene-2-ylmethyloxycarbonyl (Bsmoc) Amino-Protecting Group J. Org. Chem. 1999, 64, 4324-4338) (10/23/03 IDS Reference AR) as evidenced by Solomons (Solomons, T. W. G. Organic Chemistry Fifth Edition. New York: John Wiley and Sons. 1992, page 94, Table 3.1) and Lide (CRC Handbook of Chemistry and Physics, ed. DA Lide, 85th edn., CRC Press, Cleveland, OH, 2004-2005, web page 1) and STN (STN Express, Registry No. 141-43-5, chemical properties listing, page 1).

For *claims 28*, Carpino et al. disclose processes for the rapid solid phase and/or solution phase peptide synthesis using Bsmoc amino protecting groups in conjunction with various scavenging agents (e.g., see abstract), which anticipates claim 28. For example, Carpino et al. disclose **(a)** a coupling step, using an excess of an activated carboxylic acid component to acylate an amino component (e.g., see Carpino et al., page 4329, scheme 1 wherein H-AA<sub>1</sub>-OR is coupled to an excess of Bsmoc-AA<sub>2</sub>-OH to form Bsmoc-AA<sub>2</sub>-AA<sub>1</sub>-OR using HATU and DIEA, the excess Bsmoc-AA<sub>2</sub>-OH is removed by the NH<sub>2</sub>(CH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>N; see also page 4327, middle paragraph, “A second byproduct, derived from excess acylating agent, is the amide 16”). Carpino et al. further disclose **(b)** a quenching step in which a scavenger is used to remove residual activated carboxylic acid and also using said scavenger to deprotect the growing peptide (e.g., see page 4329, scheme 1 wherein the Bsmoc protecting group and the excess AA<sub>2</sub> are removed; see also page 4327, compounds 15 and 16). Carpino et al. disclose **(c)** the use of one or more aqueous extractions (e.g., see page 4329, scheme 1 showing removal of water soluble side products; see also page 4327, middle paragraph; see also abstract, “Application [of Bsmoc amino-protecting groups] ... represents a significant improvement over the corresponding Fmoc-based method for rapid solution synthesis due to the opportunity to use water or saturated sodium chloride solution rather than an acidic phosphate buffer to remove [i.e., extract] all byproducts”). Carpino et al. also disclose at least one step **(b)**, referred to as step **(b')**, in which an amine comprising a free anion or a latent anion is used as a scavenger of residual activated carboxylic acid (e.g., see page 4329, column 1, first paragraph wherein “ethanolamine” is disclosed). The reference does not state that

ethanolamine possesses a “free anion or latent anion”, but the Examiner contends that this would be an inherent property of ethanolamine via the following equilibrium in water  $\text{NH}_2\text{CH}_2\text{CH}_2\text{OH} \rightleftharpoons \text{NH}_2\text{CH}_2\text{CH}_2\text{O}^- + \text{H}^+$  (e.g., see Lide, web page 1, “Dissociation Constants of Organic Acids and Bases” section, Ethanolamine entry wherein  $\text{pK}_a = 12.87$ ; see also Solomons, page 94, Table 1, wherein  $\text{pK}_a$  of water = 15.74; see also STN Express registry data showing  $\text{pK}_a = 12.87$  showing that alcohol is more acidic than water, probably due to stabilization by amine). “When the PTO shows a sound basis for believing that the products of the applicant and the prior art are the same, the applicant has the burden of showing that they are not.” *In re Spada*, 911 F.2d 705, 709, 15 USPQ2d 1655, 1658 (Fed. Cir. 1990). The Office does not have the facilities to make such a comparison and the burden is on the applicants to establish the difference. See *In re Best*, 562 F.2d 1252, 195 USPQ 430 (CCPA 1977) and *Ex parte Gray*, 10 USPQ 2d 1922 1923 (PTO Bd. Pat. App. & Int.).” Finally, Carpino et al. also disclose repeating steps (a)-(c) above to synthesize a full-length peptide and/or protein (e.g., see page 4329, wherein “additional cycles” are disclosed; see also experimental section wherein longer peptides are produced).

For **claim 30**, Carpino et al. disclose pre-activation of carboxylic acid, for example, via cyanuric fluoride to produce acid fluorides (e.g., see paragraph bridging pages 4332-4333 showing activation via cyanuric fluoride; see also scheme 1 wherein HATU + DIEA is disclosed).

For **claim 31 and 36**, Carpino et al. disclose ethanolamine, which is used as a scavenger (e.g., see page 4329, column 1, paragraph 1; see also Solomons and Lide, page

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94, Table 1 showing anion).

For **claim 41**, Carpino et al. disclose one or more cycles wherein in step (b) both quenching and deprotection occur and the subsequent step (c) comprises sequential neutral extractions (e.g., see page 4320, Scheme 1, Bsmoc-AA<sub>2</sub>-AA<sub>1</sub>-OR → H-AA<sub>2</sub>-AA<sub>1</sub>-OR step; see also experimental; see also page 4327, column 2, middle paragraph, “It has now been found that the process can be simplified by switching to Bsmoc chemistry since the byproduct adduct 15 formed in this case is soluble in water, thus avoiding the need for extraction with an acidic buffer. This results in fewer complications with emulsions and loss of growing peptide into the aqueous phase.”).

For **claim 42**, Carpino et al. disclose the use of sodium chloride (e.g., see abstract, “Application to the latter methodology represents a significant improvement over the corresponding Fmoc-based method for rapid solution synthesis due to the opportunity to use water or saturated sodium chloride solution rather than an acidic phosphate buffer to remove all byproducts”).

For **claim 44**, Carpino et al. disclose the use of ethyl acetate (e.g., see generally experimental section; see also page 4332, Methods 2 and 3 see also Table I, Bsmoc-Leu-OH entry). In addition, the Examiner notes, “the general conditions of a claim are disclosed in the prior art, it is not inventive to discover the optimum or workable ranges by routine experimentation.” *In re Aller*, 220 F.2d 454, 456, 105 USPQ 233, 235 (CCPA1955). Here, the choice of solvent would be routine.

For **claims 45 and 46**, Carpino et al. disclose, for example, room temperature, which falls within 0 to 50°C (e.g., see Experimental).

For *claim 49*, Carpino et al. disclose, for example, the use of TFA to acidolytically remove the permanent protecting groups (e.g., see Scheme 1, last step).

For *claim 55*, Carpino et al. also disclose a separate deprotection step followed by one or more aqueous extractions (e.g., see Scheme 2 wherein each of the amino acids represent a “separate” deprotection step when reacted with  $N(CH_2CH_2NH_2)_3$  followed by aqueous extractions; see also Table 2 wherein Boc, T-Bu etc represent separate deprotections; see also Experimental section).

### *Response*

5. Applicant's arguments directed to the above 35 U.S.C. § 102 rejection were fully considered (and are incorporated in their entirety herein by reference) but were not deemed persuasive for the following reasons. Please note that the above rejection may be modified from its original version to more clearly address applicants' newly amended and/or added claims and/or arguments.

[1] Applicants argue, “Applicants respectfully disagree with the Examiner's assertions and conclusion and submit that independent claim 28 is not anticipated by Carpino et al. as allegedly evidenced by Solomons and Lide for the reasons stated below. Applicants assert that the  $pK_a$  value of 9.5 for ethanolamine refers to the protonation of ethanolamine (equilibrium between  $NH_3^+CH_2CH_2OH$  and  $NH_2CH_2CH_2OH$ ), while the  $pK_a$  value of 16 refers to the dissociation of ethanolamine (equilibrium between  $NH_2CH_2CH_2OH$  and  $NH_2CH_2CH_2OH$ ). The latter is indeed the equilibrium that is of importance for the DioRaSSP invention, i.e. the anion formation that plays a crucial role in the removal of quenched compounds within the protocol of



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the present invention (e.g., see 10/10/06 Response, page 5)

[1] The Examiner agrees that the 9.5 value is incorrect. The true value is ~12.87 (see newly amended rejection above). However, this does not change the fact that the alcohol is more acidic than water and, as a result, would remain to an appreciable extent in the anionic form.

[2] Applicants argue, within the quenched compounds the amine function of ethanolamine would not even be present as such; rather, at the stage of the aqueous extractions, it has been acylated by the activated carboxylic compound to yield an ethanolamide function of a considerably higher  $pK_a$  value. (e.g., see 10/10/06 Response, paragraph bridging pages 5 and 6)

[2] Applicant's arguments do not rise to the level of factual evidence. See MPEP § 716.01(c): The arguments of counsel cannot take the place of evidence in the record. *In re Schulze*, 346 F.2d 600, 602, 145 USPQ 716, 718 (CCPA 1965). Moreover, the "unreacted" ethanolamine would still function as a scavenger.

[3] Applicants argue, "Assuming that peptide synthesis would allow the application of pH values of approximately 15.74, dissociation and thus active extractive removal of quenched compounds would still be an equilibrium and therefore incomplete. However, it is also known to a person skilled in the art, that peptide synthesis is not compatible with such strongly basic (aqueous) conditions, which would result in not merely destruction of the peptide, but also of the preferred solvent of the present invention (ethyl acetate). The pH value during peptide synthesis should not exceed a pH value of approximately 12, corresponding to a dissociation extent of the ethanolamide function of 0.01 %" (e.g., see 10/10/06 Response, page 6).

[3] The Examiner respectfully disagrees. It's unclear how the 0.01% calculation has been obtained but given a  $pK_a$  value of 12.87 for ethanolamine, one would expect approximately 11%  $NH_2CH_2CH_2O^-$  at pH 12 using  $pH = pK_a - \log([A^-]/[HA])$ . This is still a significant amount of anion. Furthermore, even if, for the sake of argument, Applicants' calculations were taken as gospel (i.e., 0.01 %) it would still read on the claims (see [4] below for a more extreme example).

[4] Applicants argue, "It is further asserted that Carpino et al. fail to teach one skilled in the art that the alcohol moiety of the ethanolamide function is an anion that can be used as a scavenger. Instead, Carpino et al. teach that the alcohol moiety of the ethanolamide function remains intact during aqueous washings and does not form an anion. This assertion is underlined by the fact that extractions within the Carpino protocol are performed at neutral pH corresponding to a dissociation extent of the ethanolamide function of  $10^{-7}$  %. Accordingly, the ethanolamide function in the presence of aqueous washings as is described in Carpino et al. would not function as a scavenger as is recited" (e.g., see 10/10/06 Response, page 6).

[4] Again, Applicants assertion that only ethanolamide exists is entirely unsubstantiated (see [2] above). However, even if we assume, for the sake of argument, that only ethanolamide exists and that it only remains at a concentration of  $10^{-7}$  in its "anionic" form, that anion would still anticipate the claims because a person of skill in the art would expect that anion to scavenge just the same as any other anion albeit to an extent that is proportional to its decreased concentration. That is, Applicants' arguments are not commensurate in scope with the claims because Applicants do not set forth a "bottom limit" to the amount of scavenging that need

occur.

Accordingly, the 35 U.S.C. § 102 rejection cited above is hereby maintained.

***Claim Rejections - 35 USC § 103***

6. Claims 28-31, 36, 41, 42, 44-46, 48, 49 and 55 are rejected under 35 U.S.C. 103(a) as being unpatentable over Carpino et al. (Carpino, et al. "The 1,1-Dioxobenzo[b]thiophene-2-ylmethyloxycarbonyl (Bsmoc) Amino-Protecting Group J. Org. Chem. 1999, 64, 4324-4338) (10/23/03 IDS Reference AR) and Tolle et al. (WO 00/71569) (Published November 30, 2000) and Houghten et al. (Houghten, R.A.; Pinilla, C.; Blondelle, S.E.; Appel, J.R.; Dooley, C.T.; Cuervo, J.H. "Generation and use of synthetic peptide combinatorial libraries for basic research and drug discovery" *Nature* 1991, 354, 84-86).

For *claims 28, 30, 31, 36, 41, 42, 44-46, 49 and 55*, Carpino et al. teach all the limitations stated in the 35 U.S.C. 102(b) rejection above (incorporated in its entirety herein by reference), which anticipates and, as a result, renders obvious claims 28, 30, 31, 36, 41, 42, 44-46, 49 and 55.

The prior art teaching of Carpino et al. differ from the claimed invention as follows:

For *claim 29*, the prior art teachings of Carpino et al. differ from the claimed invention by not specifically reciting the amounts of reagents as carboxylic component, coupling additive greater than coupling reagent greater than amino component. Carpino et al. only show carboxylic component greater than amino component.

For *claim 48*, Carpino et al. fail to teach the use of automation.

However, the combined references of Tolle et al. and Houghten et al. teach the following limitations that are deficient in Carpino et al.:

For **claim 29**, the combined teachings of Tolle et al. and Houghten et al. teach the use of coupling reagents used in conjunction with coupling additives (e.g., see Tolle et al., page 11, line 3 wherein activated N-hydroxysuccinimide esters are used; see also Houghten et al., Tables and figure). In addition, differences in concentration (e.g., carboxylic component, coupling additive greater than coupling reagent greater than amino component) will not support the patentability of subject matter encompassed by the prior art unless there is evidence indicating such concentration is critical. “[W]here the general conditions of a claim are disclosed in the prior art, it is not inventive to discover the optimum or workable ranges by routine experimentation.” In re Aller, 220 F.2d 454, 456, 105 USPQ 233, 235 (CCPA 1955). Here, it would be conventional and within the skill of the art to *identify the optimal concentration*. It is well-established that merely selecting proportions and ranges is not patentable absent a showing of criticality. In re Becket, 33 U.S.P.Q. 33 (C.C.P.A. 1937). In re Russell, 439 F. 2d 1228, 169 U.S.P.Q. 426 (C.C.P.A. 1971).

For **claim 48**, the combined teachings of Tolle et al. and Houghten et al. disclose the use of automation (e.g., see Tolle et al. figure 2).

It would have been obvious to one skilled in the art at the time the invention was made to use the scavenging resins for the combinatorial synthesis of proteins as taught by the combined teachings of Tolle et al. and Houghten et al. with the Bsmoc Amino protecting groups as taught by Carpino et al. because Carpino et al. state that their

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method can be used with scavenging resins and that it is also applied to combinatorial synthesis i.e., the references represent analogous art. Furthermore, one of ordinary skill in the art would have been motivated to use the scavenging resins as taught by Tolle et al. because Tolle et al. explicitly state that their resins will “minimize the requirement for isolation of intermediates” that are produced in peptide synthesis using scavengers (see Carpino et al., Field of the invention), which would encompass the peptide synthesis disclosed by Carpino et al. In addition, Houghten et al. teach that their “split and mix” method can be advantageously used to produce large peptide libraries (e.g., see Houghten et al., abstract), which would encompass the peptide libraries of Carpino et al. Finally, one of ordinary skill in the art would have reasonably expected to be successful because all three references teach the successful synthesis of peptides and both Carpino et al. and Tolle et al. teach successful examples of using amine scavengers in peptide synthesis.

### *Response*

7. Applicant’s arguments directed to the above 35 U.S.C. § 103(a) rejection were fully considered (and are incorporated in their entirety herein by reference) but were not deemed persuasive for the following reasons. Please note that the above rejection has been modified from its original version to more clearly address applicants’ newly amended and/or added claims and/or arguments.

[1] Applicants argue, “[t]he arguments proffered above to address the §102 rejection apply equally well to this rejection, namely that at such high pH values, i.e., pH 16, peptide synthesis would not be compatible with such strongly basic conditions and would result in the

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destruction of the peptide. Further, extractions at a neutral pH as is described in Carpino et al. would correspond to a dissociation extent of the ethanolamide function of  $10^{-7}$  % and thus the alcohol moiety of ethanolamide function would remain intact following the protocol of Carpino et al. Accordingly, the ethanolamide function subjected to aqueous washings at a neutral pH as described by Carpino et al. would not function as a scavenger as recited in claim 28. In addition, neither Tolle et al. or Houghten et al. describe an amine comprising a free anion or latent anion that could function as a scavenger" (e.g., see 10/10/06 Response, page 6, last 2 paragraphs).

[1] To the extent that Applicants are merely repeating their previous arguments, the Examiner contends that those arguments were adequately addressed in the above sections, which are incorporated in their entireties herein by reference. Thus, Tolle et al. and/or Houghten et al. are not required to remedy these alleged deficiencies.

Accordingly, the 35 U.S.C. § 103(a) rejection cited above is hereby maintained.

### ***Double Patenting***

8. Claims 28-39, 41, 42, 44-46, 48, 49, 51, 52 and 55 are provisionally rejected under the judicially created doctrine of obviousness-type double patenting as being unpatentable over claims 28-51 of copending Application No. 10/692,354 (2004/0082760 A1) (referred to herein as '354). An obviousness-type double patenting rejection is appropriate where the conflicting claims are not identical, but an examined application claim not is patentably distinct from the reference claim(s) because the examined claim is either anticipated by, or would have been obvious over, the reference claim(s). See, e.g., *In re Berg*, 140 F.3d 1428, 46 USPQ2d 1226 (Fed. Cir. 1998); *In re Goodman*, 11 F.3d 1046, 29 USPQ2d 2010 (Fed. Cir. 1993); *In re Longi*,

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759 F.2d 887, 225 USPQ 645 (Fed. Cir. 1986). Although the conflicting claims are not identical, they are not patentably distinct from each other because, for example, claims 28-39, 41, 42, 44-46, 48, 49 and 51-55 are generic to all that is recited in claims 28-51 of '354 or represent overlapping scope.

For *claim 28*, the '354 application discloses the same method steps (a)-(c) and also the same method step (b)' and thus is identical to the claimed method with the exception of the optional step (d) recited in the '354 application.

For *claims 29-39, 44-46, 48, 49, 51 and 52*, the '354 application discloses the exact same method steps (e.g., see '354 application, claims 29-39, 41-44, 46-48, 50 and 51).

For *claims 41 and 42*, the '354 application discloses that both quenching and deprotection can occur (e.g., see '354 application, claim 28). In addition, the "basic" extractions disclosed in claim 41 of '354 anticipate claim 41 of the present invention. Furthermore, it would have been obvious to one having ordinary skill in the art to modify embodiments of '354 that fall outside the scope of the present application (e.g., the acidic extractions) to select a specifically disclosed embodiment that falls within the scope of the present application (e.g., the basic or neutral extractions) because these embodiments describe similar method steps (e.g., extraction) with similar results (e.g., purification). One having ordinary skill in the art would have been motivated to do this because these embodiments (e.g., neutral and basic extractions) are disclosed as being preferred embodiments in the '354 application and the dependent claims of '354 teach toward Applicants' claimed invention (e.g., see claims 42 and 43 of the '354 application).

For *claim 55*, the '354 application also describes a separate deprotection step (d),

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followed by one or more aqueous extractions (e.g., see '354 application, claim 28, step (d)).

This is a provisional obviousness-type double patenting rejection because the conflicting claims have not in fact been patented.

### *Response*

9. Applicant's arguments directed to the above double patenting rejection were fully considered but were not deemed persuasive for the following reasons. Please note that the above rejection has been modified from its original version to more clearly address applicants' newly amended and/or added claims and/or arguments.

[1] Applicants argue, "In response, Applicants will address the obviousness-type double patenting rejection upon indication that claim 28 is deemed to be allowable except for the obviousness-type double patenting rejection." (e.g., see 10/10/06 Response, page 7).

[1] The rejection will not be held in abeyance (e.g., see MPEP § 804 B. Between Copending Applications—Provisional Rejections, "The 'provisional' double patenting rejection should continue to be made by the examiner in each application as long as there are conflicting claims in more than one application unless that "provisional" double patenting rejection is the only rejection remaining in one of the applications."). Here, a double patenting rejection is NOT the only rejection remaining in one of the applications and thus the double patenting rejection is proper.

Accordingly, the double patenting rejections cited above are hereby maintained.



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***Conclusion***

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Jon D Epperson whose telephone number is (571) 272-0808. The examiner can normally be reached Monday-Friday from 9:00 to 5:30.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, James (Doug) Schultz can be reached on (571) 272-0763. The fax phone number for the organization where this application or proceeding is assigned is (571) 273-8300.

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to the receptionist whose telephone number is (571) 272-1600.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

Jon D. Epperson, Ph.D.  
January 21, 2007

JON EPPERSON  
PRIMARY EXAMINER

